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                            to the CTV - Frequencies of amin -substituted beta -last an autibiotics
                              orrrises enzyme-paralyseo reaction i amin - keta -lastams and
                              amino-substituted anylated compounds, employing solubilization with an
                              acid and treatment with a buffer to regelerate the enzyme.
                                                      DETAILED DESCRIPTION - Preparation of amino-substituted beta -lactam
                             antibiotics comprises reacting the corresponding amino- beta -lactam
                              components and amino-substituted abylated side chain forming compounds
                             using a covalently immobilized enzyme datalyst. The reaction
                            is objects are solublished at a pH ^{\circ} Tol. Candia temperature of ^{\circ} -5 degree .
                            the presence of the enzyme, which is about a color a general mapping as a little enzyme as
                             separated oir and regenerated by treating with a neutral butter at ak W
                             dea. C.
                                                          USE - The technique is useful for the preparation of:
                                                               ampicillin from 6-aminopenicillanic
                              acid 6-APA .... D-
                            and D-phenylglycine retay, estated
                             [-}::01:y1.51:y5:1:37...3=/
                                                             go objaklom irom s-mila (= 1-ANIA and D-
                            phenylglycine methyl ester or l-phenylalyalnamide;
                                                   4° amexicillin iron 6-APA and
                              D-4-hydrowymichyl glycino mothyl oster in 1-4-hydr wymienyl flycinami ie;
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Enzymatic propa . In the area of the transfer and and it is to be, an ampicilling a second of INHAH, E; BEARARE, omir-m omirar rimya danayot ur it arri ad BEFORE VARY AND TRAKELOT OF THE SECRET THE SECRET TO DESIGN AND MESSAGE AND SECRET THE SECRET THE SECRET AND SECRET THE SECRET TO DESIGN AND MESSAGE AND SECRET THE S # 7F 11 ANSWER 1 OF 6 WEIDS CUPYRIGHT 2000 DERMENT INFORMATION DID NO 9918832 A TRABE 19990813 NOTENTY - Resovery of Beta -lastam and Blotin from mixture sontaining and the transfer D-phenylglycine in solution is new. ល់ស្រួល ស្រួស៊ីសម្រើស៊ីស៊ីទី អ៊ីស្នាក់ កាន់ស្លាក់ សុទ្ធសម្រេង នេះ និងស្រួសស្រាស់ ស្រួសស្រួសស្រួស សុទ្ធសម្រាក់ សុទ្ធសម្រេង ប៉ាក់ស្រួសស្រួស ប៉ុន្តែ និងសុ % n %moration sim that D-phenylglycine remains in 3.144 1.11; (b) recovering the solid beta -lactam antibiotic; (c) subjecting the remaining liquid to temperature in wease to 1 med deg. 3 with formation of solid D-phenylglycine; ij separating out the D-phenylglycine as solid; e; partially resinculating the mother liquor. Activity - Antibilitie. TGE - Used to recover approx. -b lastam antibiotics from mixtures containing antibiotic and D-phenylglycine (claimed). May be used to recover antibiotics with a phenylglycine side-shain e.g. sefalexin, ampicillin, sefactor, pivampicillin, becampicillin, talampicillin and defaloglycine. An enzyme reactor (1.8); 11-cm diameter; fitted with a 178 approx. nom mesh sieve better was filled with 300 z zer tot. bottom was filled with 300 g net-wet Assemblase (RTM; immobilized Escherichia coli penicillin acylase from 8. coli ATCC 1105). A preparation. pearture l.v. it was colled with 6-APA lelve g',
D-phenylglycine arise of log and water 4 ml at log and value. The stories on at the . The temperature was heli at 1 des. 0. D-phenylglycine amide sulfate solution (42). The was added at a constant rate over 138 minutes. The pH was approximately equal (.). From = 198 minutes onwards, the pH was maintained at 6.3 by titration with (II sulfuric acid. At 5t = 570 minutes, the amount of ampicillin was numinum and the pH reduced to 4.7 by addition of 4%sulfuric acid. enzyme ampicillin b-phenylglycine ampicillin D-phenylglycine ampicillin D-phenylglycine phenylglycine slurry was removed from the enzyme reaction phenyigiyoine stati, assismited tith the enzyme leaster wis the sleve bottom by means of stirred tiltration using a pitched-blade stirred positioned 1.8 cm above the sleve. Stirring was in an upward direction at approximately equal of type. The slurry separate if no the leaster was filtered on a 38 class filter. The wet cake was put asib and the mother liquor returned to the enzyme reactor. Stirred the morner liptor resulted to the enzyme teactor, othered cities for followed by B filtration of the slurry resumed. The enzyme result of was washed with ampiculin-D-phenylglycine of the bound of the liptor of liptor of last of the days of rothed with the followed with

war age to the ampicillin D-phenylglycine

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               -phenylglycine prijusei in the enzyme reast i.
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               APA and of g D-phenylglycine.
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                NOVERLY - Bus very or beta -lasten entiristic from mixture containing
                 antibuotis and D-phenylglycine in solution is new.
                               peramen međem dibli - Bar very comprisasi
                                (a) bringing mixture to pH 3-3 at neg. (-2) des. 2 at concents click
                such that D-phenylglycine remains in solution;
                                 (b) recovering solid beta -lactam antibiotic;
                                  o' subjecting remaining liquid to concentration step in which
 slurry
                with solid beta -lastam antibioris and solid D-
                phenylglycine is well; 87
                                 A Teach anna scharge to the second school and do the first of the
                  annication dies lyes:
                                e separating D-phenylglycine as solid and;
                                  to ลาไปปละ ผู้ละประชาที่ ซึ่งปีประเทศ กาละ Nets -lantam antibi การการทะสะบา
                 in the mother liquin.
                               ACTIVITY = Actual Color,
                               USE - Used to recover approx. -b lastam antibiotics from mixtures
                 Some aiming and iming the ani D-phenylglycine claimed.
                Marchandsein, persyar antibiotics with a phenylglycine
                 sike manneth een verstekin, ampicillin, verstek r, plaas i villin,
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                 rearror 1.0 10 was filled with 6-APA (181.6 %)
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. The temperature was held at 1 tem. 7. Dephenylglycine and temperature solutions 40%, 7% to was added at a
       To not and Transcover is to win where I have pH was rapportunately repeal for the body
      which provides cowards, the pH was raintained at 0.5~{\rm ky} titration with 0.0
      sulfuric acid. At it a 10, minutes, the amount of ampiculin was naminum and the pH reduced to 4.0 by addition of all
      phenylglycine visity was terrived from the enzyme resultat
      Fig The sieve postom by means of sti ried filtration using a
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      stirrer positioned 0.8 cm above the sieve. Stirring was in an upward
      diresti h an approximately equal 511 rpm. The reaction was flushed with water 11 mountably to mile at 1 deg. The wash waters were also
      that the likewise the many letters a first rate by the clarry and also waste
      were compared at 1 deg. ). The resulting slurry contained at \tau of the total amount of ampicillin and greater than 48.8 of the total
      amount of D-phenylglycine produced in the
       enzyme reactor. A mother liquor from an ampicillin
      orystallization step of Mb g; 1 deg. 0; pH 6.11 Juntained 6.36 6 -APA, 0.11 D-phenylglycine amide, 1.53
       ampicillin and 0.98 [D-phonygloying. The nother liquor was our entraced by evaporation with the aid of a TFE. The feed was supplied
       from a storage vessel and the product returned to the same storage
vessel,
      which was stirred. The wall temperature of the TFE was adjusted to 65
dea.
       C and the pressure to 80 mbar. Circulation started at t = 0. The liquid
       temperature was maintained at 40 deg. C. Evaporation was stopped at t \sim
       257 minutes, by which time 5655 g concentrate had been collected. The
       contents of the T FE and storage vessel were diroulated for a further (
       contents of the Tork and storage vessel were circulated for a further 1 hours while the temperature was 1 wered linearly from 41 deg. Total wat. The The non-versage vessel were thank i with 10 ml water than 10 ml water at 10 deg. The tendensate and 1st wash water were confined and filtered this glass filter (1-cm diameter; 10 minutes). The solid was re-washed with water (25 ml) at 1 minutes.
       dea. Compaire ampicillin D-phenylglycine
       wei cake will go. The mother liquer was discharged. The wet cake was quantitatively transferred to a stirred reactor with water (40% g). The
       slurry was stirred for I hour at 1 deg. 7. Hydrothloric asid (60) 33 g was added over 4 minutes at 8 deg. 7. Attends think for a number, the
       slurry was filtered on a glass filter (13- m diameter; f minute) and the D-phenylglycine was take washed with water (10 ml at 15 ml and the D-phenylglycine (10 ml at 15 ml).
       mother liguor and wash water were added to acid solution prepared by
       reorystallization of ampicil lin. The final yield was difampicillin. 3880 including nuclei or 100 a ampicillin
       .3H22 excluding nuclei siving 92 ampicillin.3H22 relative to
            mmol 6-APA.
             ADVANTATE - Fromess is simple and ran be applied in an investment
       scale. Itsses of port -last am antibulting as stilling reduced and
       D-phenylglycine is to reverse. D-
       phenylglycine can be recovered selectively enabling complete
        indulus និងសម្រាប់សម្រាប់ប្រជាសេស អាសាសាណភាពល និងសាធិនី - នៃការយោធារាអំពេញ ១០
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                   Recovery of approximatelybeta-lastam antibiotic from solution of
                    antibilitie and D-phenylglycine.
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                          The preparation of ampicillin in which 6-
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                  aminopenicillanic acid 6-APA is
                         subjected to an enzymatic adylation reaction with a theny!
                          glyrine derivative such as phenylglycine is claimed. The
                 reaction is instantenized by the total concentration of 6- APA and ampicillin together being below 390 mM. The
                         molar ratio of adylation agent to 6-APA is below 2.0.
                          By ensuring the concentration of 6-APA in the
                         dissolved form present in the reaction mixture is kept relatively low, a
                          higher conversion can be achieved. The stirrability of the reaction
                         mixture is also considerably improved. The phenylglycine
                          derivative is metered in as a salt of phenylglycine amide (PGA)
                  and an avide, p. as a solution of D-FGA/O.E sulfuric acid in water. The pH of the reaction mixture is lowered as soon
                          as near-to-maximum conversion is achieved. [20pp]
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                           Preparation of ampicillin with higher nerversion and yield;
                                    by enzyme adylation of 6-aminopenicillanic
                               acid with a phenyl glycine derivative e.g.
                              phenylglycine
                           Moody H M; Boetsen W H J
   ĒĀ
                           Heerlen, The Netherlands.
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                         phenylglycine, and beta-last and derive are sept out. After the
                         enzymatic reaction, the obtd. mixt. from which at least the
                         enzyme and solid D-phenylglycine have been
                         removest, is treated with an aldebyte at a pH of 7.8-+.8 and the Schitt
                         base of D-phenylglycine and he is sepai. Out.
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                    ាការសម្នាធាតិ សមិស៊ី រ៉ឺនការសេស សាសសការការការជា ៖១ ជាធាកាសភា
    s limiting if the latter is higher, recovery and purific of this product
    and we network, is simplified. He covery and tecycle if the D^\perp
    phenylglycine arise derived by simple filtration or extn., in
     ກໍ່ສະເນົ້າໃນຊື້ ເລກ ເປັນປະທູນິດ Benzaldungur) ເຮັນສອນ, ອະດາຍສະ ໝົນປະທູນິດ ເພນ ສອນທົດ ພະ
     extrastivent, alth ligh other solvents, or mixts with FhOHl, can be used.
     The Johnif pase is then split with acid, eg H2SO4, and resysted.
     As an excess of amide is used in the coupling to obtain a high yield or
    per selaman producer very of the exposs is necessary in order to provide
     commendately aftractive process, epitheless the mixtorypically contains
     1- belief i D-phenylglycine and be, .1-. equivs i D-phenylglycine and ...-1 - pure i dather,
     all per molecul produ
     Owş..
     1998-078240 (11)
01998-033800
     Be obvery of Depremylealy sine amide from antibiosis coupling - by
    41.20.14
     or pure, easily sepa, remyclable Schiff hase, used in enzymatic
     preprior paphalemon, betailtr, ampicillin, etc...
     POZÍBUS DIS
BOESTEN, W H J; MOGUY, H M; MOCDY, H
      (STAM) DSM NV
ΞA
                     A1 19950202 (199510)* EN 12p
     WO 9503420
         RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL ET SE
         W: CN JE KR US
         1007296 A3 19950509 (199524)
012443 A1 19970525 (199525)
          B: AB IB EN EB BE IT ML ET
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     EG LITULE: 03 1786.LIT (1986)1
WO 8503470 AT WO 1984-MINET 18840718; BE 1.07186 AR BE 1888-750 18880717;
BE 71843 AT EF 1884-814421 18840718, WO 1884-MINET 18840718; TE 11878 ST
      010 1994-190418 1994 (1996 EE TICA43 BI EE IMMA-904401 19941019) WO
1994-101081 1994 7197 1E GRAST AR E DE 1994-6 7049 IMMAINIS, EE
FP 11244 At baset of MOTER 04. ; he clear by mased on MC 1012420; DR 04417148 E Based on EF 712443, Based on MC 9873420; ES 2112286 T3 Based
      EF "12443
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                         4,4437713
       ANOMER \epsilon of \epsilon filtering correspond to definent information into
      The first in or a location at a section which a location and in the de-
       Page 1 Dephenylglycine and be the di-
     enzymatic : a with is claimed. The sold: D-
     phenylglycine and beta-last amovernments and the Johirt base are
       segurated out from the enzyme by theatment with an aldelyde at
       pH 0.144. The D-phenylglycine amide derivative is
       easily separated out in pure form, pretendily before the beta-last amprofits, as the solubility of the laster is higher. We overy and
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if the D-pheny voice anide derivative is by specification or a faction. If pencaldehyde is unlessed an extraction silvent, although other solvents, in cixtures with the search as an extraction silvent, although other solvents, in cixtures with the search as a contract of the specific pencal search in the search of the pencal search in the search of the pencal search of the sear
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